

In the Claims

1-61 (canceled).

62 (previously presented). An antibody that induces superagonistic signaling by a cell surface receptor, wherein said antibody binds to the extracellular portion of the receptor at a membrane proximal region and said receptor comprises a cytoplasmic domain which is dependent on an extrinsic protein kinase, wherein said antibody does not bind only the C'-D loop of human CD28.

63 (previously presented). The antibody according to claim 62, wherein said antibody:

- (i) binds orthogonally to the main axis of the domain of the receptor which it is binding, and/or
- (ii) which lies parallel to the cell surface when bound to the receptor, and/or
- (iii) which binds to a β -strand polypeptide chain of the receptor, and/or
- (iv) which binds within 75Å of the cell surface.

64 (previously presented). The antibody according to claim 62, wherein said receptor

- (i) comprises an ITAM motif, ITIM motif or "switch" signaling motif, and/or
- (ii) is a member of the CD28 family of proteins, and/or
- (iii) is expressed on the surface of a cell of the immune system, and/or
- (iv) comprises a cytoplasmic domain capable of being phosphorylated by a Src kinase, and/or
- (v) comprises a cytoplasmic domain capable of being dephosphorylated by CD45, and/or CD148, and/or another large receptor tyrosine phosphatase, and/or
- (vi) is one of the receptors listed in Table 2.

65 (previously presented). The antibody according to claim 62, wherein said cell surface receptor is CD28, CTLA-4, ICOS, PD-1 or BTLA.

66 (currently amended). The antibody according to claim 62, wherein said antibody binds to an epitope comprising SEQ ID NO: 23, SEQ ID NO: 24, SEQ ID NO: 25, SEQ ID NO: 26, SEQ ID NO: 27, SEQ ID NO: 28, SEQ ID NO: 29, SEQ ID NO: 30, SEQ ID NO: 31, SEQ ID NO: 32, SEQ ID NO: 33, SEQ ID NO: 34, SEQ ID NO: 35, SEQ ID NO: 36, SEQ ID NO: 37, SEQ ID NO: 38, SEQ ID NO: 39, SEQ ID NO: 40, SEQ ID NO: 41, SEQ ID NO: 42, SEQ ID NO: 43, SEQ ID NO: 44, SEQ ID NO: 45, SEQ ID NO: 46, SEQ ID NO: 47, SEQ ID NO: 48, SEQ ID NO: 49, SEQ ID NO: 50, SEQ ID NO: 51, SEQ ID NO: 52, SEQ ID NO: 53, SEQ ID NO: 54, SEQ ID NO: 55, SEQ ID NO: 56, SEQ ID NO: 57, SEQ ID NO: 58, SEQ ID NO: 59, SEQ ID NO: 60 or SEQ ID NO: 61, selected from:

<i>Protein</i>							
hPD-1	PALLVV;	DNATF;	RMSPSNQTDK;	QPGQDCRF;	MSVVR;	NDSCFY;	LRAELR;
hBTLA	QSEHSI;	DPFEL;	KLNG;	QTSWK;	LHPFP;	NDNGSY;	TTLYVT;

<i>Protein</i>							
hCD28	SPMLV;	AVNLS;	SLHKGLDSAVEVCV;	VYSKTGFNCDO;	FYLQN;	TDIYFC;	NGTIHV;
hCTLA-4	PAVVL;	GIASFV;	TVLRQADSQVTEVCA;	FLDDSICTG;	LTIQG;	TGLYIC;	NGTQIVV;
hICOS	YEMFI;	GVQIL;	QLLKGGQILCD;	VSIKSLKFCHS;	FFLYN;	ANYYFC;	TGGYLHI;

<i>Protein</i>	
hCD28	GNYSQQQLQVYSKTGF;
hCTLA-4	YMMGNELTFLDDDS;
hICOS	KTKGSGNTVSIKSLK; or
hPD-1	LAAFPEDRSQPGQDCR;